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PRINCIPAL INVESTIGATOR:

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13. SUPPLEMENTARY NOTES

14. ABSTRACT

The HBCU Summer Research Training Program accepted a total of 17 students from Lincoln University for the eight week session during the summers of 2010, 2011, and 2012. Each student was assigned to a laboratory of a participating mentor and also paired with a member of the mentor's laboratory. This laboratory member assisted with day to day aspects of the research project. During the summer the students worked diligently on their research project, participated in meetings of the mentor's laboratory, attended workshops and seminars associated with our and other summer programs, and attended a special course in prostate cancer. We integrated the Lincoln students into social programs held throughout the campus for summer interns and they attended and participated in the CIC Conferences. At the end of the summer sessions the students presented a poster of the research results from the summer experience. They also presented the results of their research in the fall at Lincoln University. Of the students that have graduated from Lincoln, approximately 72.4% are attending postgraduate programs.

15. SUBJECT TERMS

Summer research; prostate cancer; HBCU institutions

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Introduction:

In our initial award (W81XWH-06-1-0266), begun in 2006, we were funded for five students from Lincoln University of Pennsylvania. Because of a large number of qualified student applicants we were funded for additional three students in 2007 (W81XWH-07-1-0241), allowing our program to support a total of eight Lincoln students each summer. We applied for, and were awarded; two additional grants (W81XWH-09-1-0270) and this award (W81XWH-10-1-0459) after the original grants had been completed. For the year reported here we had the following faculty participants: David M. Lubaroff, PhD, Principal Investigator, Paul Heidger, PhD, University of Iowa Faculty Advisor, Derek Swinton, PhD, Lincoln University Faculty Advisor, and the following mentors: Jackie Bickenbach, PhD, Andrean Burnett-Simons, PhD, Elizabeth Chrischilles, PhD, Frederick Domann, PhD, Elizabeth Field, MD, Michael Henry, PhD; Siegfried Janz, MD, Yi Luo, MD, PhD, Aliasger Salem, PhD, Michael Schultz, PhD, Douglas Spitz, PhD, George Weiner, MD, Michael Wright, PhD, and Nicholas Zavazava, MD.

Body:

Recruitment and Admission:

Brochures, application forms, and posters were designed and printed and sent to Dr. Swinton at Lincoln and one of the faculty mentors PI traveled to Lincoln University in January 2010, January 2011, and January 2012, met with Dr. Swinton, Dr. John Chikwem, and the Chairmen of the Departments of Chemistry and Biology. Presentations were made about the summer training program to groups of students at the beginning of their classes. Nineteen applications were received for the 2010 summer session, 16 for the 2011 summer session, and 16 for the summer of 2012. The applications were reviewed by the Admissions Committee whose membership consisted of Dr. Lubaroff, Dr. Heidger, Dr. Henry, Dr. Domann, and Dr. Swinton. After making offers to students admission was offered to a total of ten students for the summer of 2010 and eight students for the summer of 2011, and eight students for the summer of 2012. In 2011 one of the students retracted their acceptance for personal reasons.

Students Participating in the 2010 Program:

Theresa Akede Christiana Awoyemi Kaylene Baugh Nakita Brown Chalwe Diallo Danielle Holsey Jhanelle Marks Danielle McKnight Stephanie Rand Stephen Sangster

Students Accepted for the 2011 Program

Jhoneil Cooper
Darah Doubt-Swinton
Jodi-Ann Foster
Patrick Ihejirika
Candice Lynch
Ayanna Raeburn
Nathaniel Sangster

Students Accepted for the 2012 Program

Laurie-Ann Davis
Chalwe Diallo
Shauna Ebanks
Ashley Ellis
Shakeema Jones
Candice Lynch
Jhanelle Markes
Rasheid Smith

Advance Preparation and Information Distribution:

Following acceptance of the students into the program we assigned each student a mentor based upon his/her choices listed in their applications. Each mentor then assigned a member of the lab as a "big brother/big sister," a person that partners with the student during the 8 week summer session. The mentor also prepared a portfolio of articles covering the area of research the student would be working on, including published papers by the mentor. These materials were sent to the students in advance of their arrival at the University of lowa.

A six week course on Prostate Cancer was organized with six faculty assigned to deliver lectures. The following represents the course schedule with lecturers:

Iowa-Lincoln Summer Research Training Program - 2011 Prostate Cancer Course Room 3240 MERF

Lecture	Date	Subject	Lecturer
Week 1	June 14	Introduction to cancer	Spitz
Week 2	June 21	Basic aspects of prostate cancer	Dahmoush
Week 3	June 28	Epidemiology of prostate cancer Singh	
Week 4	July 5	Genetics of prostate cancer	Domann
Week 5	July 12	Clinical treatment of prostate cancer	Vaena
Week 6	July 19	Immunotherapy of prostate cancer Lubaroff	

Iowa-Lincoln Summer Research Training Program - 2012 Prostate Cancer Course Room 2156 MERF

Lecture	Date	Subject	Lecturer
Week 1	June 12	Introduction to cancer	Spitz
Week 2	June 19	Basic aspects of prostate cancer	Dahmoush
Week 3	June 26	Epidemiology of prostate cancer	Gupta
Week 4	, July 3	Genetics of prostate cancer	Domann
Week 5	July 11	Clinical treatment of prostate cancer	Vaena
Week 6	July 17	Immunotherapy of prostate cancer	Lubaroff

Housing and meal plans were arranged in collaboration with the Iowa Biosciences Advantage Program (IBA). Lincoln students were paired with IBA students in the dormitory. Plans were also formulated to integrate workshops, lectures, and social events with other programs dedicated to the training of minority students, such as the Iowa Alliance for Graduate Education and Professiorate (AGEP), CIC Iowa Summer Research Opportunities Program (SROP), and Iowa Biosciences Advantage (IBA) Program.

The Summer Program:

A welcoming summer picnic was held on the day of the student's arrival in Iowa City in conjunction with the other summer programs at the University of Iowa. The following day the students wet with the PI, administrator, mentors, and big brothers/sisters for an orientation and then taken to the laboratory of their mentors to begin the summer research training program.

During the 8 w eek session each student worked diligently with his/her mentor and I ab partner on the assigned research project. Each student had an independent project. They attended seminars, workshops, lab meetings, journal clubs and the weekly lectures in the program's prostate cancer course (see schedule above). During the seventh week the PI met individually with each student to evaluate his/her summer experience. The unanimous opinion was that the program was a success. The students indicated that they learned much about research, about prostate cancer, and about the advantages of a research career. At the end of the summer session each student presented their research as part of a poster session held during an afternoon of the last week. In addition to the poster presentations each student gave an oral summary of their research project to the mentors, big brothers/sisters, and other summer students. In addition to the mentoring the students received from their research lab and the PI, they received career counseling. We discussed the options for each of them based upon their experience and their desire for the type of future they envisioned for themselves. Among the topics discussed was graduate school versus medical school, their ultimate goals of research, patient care, and/or teaching.

All of the students attended the CIC Conference held July 23-25, 2010 at the Ohio State University, Columbus, OH. E ach student presented a pos ter of their research and participated in roundtable discussions with students from other CIC-associated institutions. Dr. Heidger accompanied the students on the trip. The students did not attend the 2011 conference since it was scheduled for an earlier date and there would have been insufficient time for their research to yield significant results.

Follow-Up:

We made frequent contact with all of the students after their departure from the University of Iowa. The mentors, faculty advisors (Heidger and Swinton), and mentors, all had contact with the students since the end of the 2006 summer session. Many of the mentors were asked to write letters of reference for the students' applications to graduate schools. The following table reports on the current status of the 2006, 2007, 2008, 2009, 2010, and 2011 summer students.

Lincoln Student Follow-Up

Name	Year	School	Program or Current Year at Lincoln	
Oluwaseun Adekanye	2006	U. Michigan	Obtained MD from Penn State Coll. Of Med., currently a resident at Michigan.	
Shaynah Browne	2006	U. Mass	graduated with MS; working in lab at Albert Einstein Med Ctr, NY	
Nikesha Haynes	2006	U. Rochester	graduate school	
Shivaughn Johnson	2006	Ross University Medical School	medical school; left and working	
Briquel Sherman	2006	University of West Indies	medical school	
Shaan Spence	2006	U. South Florida	graduate school	
Bisola Awoyemi	2007	Univ. of the District of Columbia	obtained MS degree, currently working in lab at Harvard.	
Seme Diallo	2007	Drexel University	obtained MS degree; currently working studying for MCAT to enter medical school	
Caroline Dias	2007	none at this time	working & applying to grad schools	
Titilope Idowu	2007	Morehouse College	graduate school (public health)	
Patrick Ndungu	2007	University of Iowa	graduate school	
Elizabeth Okyne	2007	U. Iowa		
•			nursing school	
Katrina Probherbs	2007	Adelphi University	graduate school	
Bukola Fatunmbi	2008	U. Mass	graduate school	
Katherine Foster	2008	Fox Chase Cancer Center	working in research lab	
Theon Francis	2008	Healthcare facility	Government assisted training prog. in lab. science	
Michelle Gray	2008	Johns Hopkins	working in laboratory	
Julia Greenfield	2008	U. Maryland	graduate school	
Gladys Murage	2008	U. Mass	graduate school	
Brittany Stokes	2008	none at this time	working in health care & applying to grad. Sch.	
Stacy-Ann Wright	2008	none at this time	working at Fox Chase Cancer Center	
Kaylene Baugh	2009	U. Pennsylvania	research internship	
Christina Chisolm	2009	U. Mass	graduate school	
Seme Diallo	2009	see 2007	see 2007	
Elizabeth Okyne	2009	see 2007	see 2007	
Stephen Sangster	2009	none at this time	teaching at college level; applying to graduate school	
Keyana Tyree	2009	none at this time	obtained MS degree from U. Mass; currently working at Fox Chase CC; has applied for PhD program	
Neja White	2000	none at this time	working at local hospital and apply for medical school	
Akede, Theresa	2009		graduata	
,	2010	U. Maryland Baltimore	graduate	
Awoyemi, Christiana	2010	Cameron	senior; plans to apply to medical school	
Sangster, Stephen	2010	see 2009	see 2009	
Rand, Stephanie	2010	Thomas Jefferson	medical school	
McKnight, Danielle	2010	Lincoln	senior	
Markes, Jhanelle	2010	Lincoln	senior	
Holsey, Danielle	2010	Lincoln	senior	
Diallo, Chalwe	2010	Lincoln	junior	
Brown, Nakita	2010	U. Pittsburgh	postbaccalaureate program	
Baugh, Kaylene	2010	see 2009	see 2009	
Cooper, Jhoneil	2011	Lincoln	senior	
Doubt-Swinton, Darah	2011	Lincoln	junior	
Foster, Jodi-Ann	2011	Lincoln	senior	
Ihejirika, Patrick	2011	Lincoln	junior	
Lynch, Candice	2011	Lincoln	senior	
Raeburn, Ayanna	2011	Lincoln	junior	
Sangster, Nathaniel	2011	Lincoln	junior	

As is evident from the table, of the students that have graduated from Lincoln, 21 of the 29 (72.4%) are attending postgraduate programs (post-baccalaureate, graduate or medical). An additional 5 or 17.2% have plans to apply for postgraduate programs. If these 5 students are successful in entering these programs we will have total of 88.4% of the graduated students continuing their education. This is an amazing statistic. On e of our former students, Ms. Nikesha Haynes, was highlighted at the 2011 IMPaCT conference as evidence of the success of the HBCU Summer Research Training Program in Prostate Cancer. She is a graduate student at the University of Rochester.

Key Research Accomplishments

Each of the students worked on research projects that were part of an overall program within the laboratory of their mentors. As such, it is difficult to identify key research accomplishments for each student research project. Continuation of the research program by each mentor will certainly produce important research findings, aided in part by the summer research of the Lincoln University students. What is key is the mentoring and counseling of the students to aid in their future as scientists in the area of prostate cancer research. The high percentage of the students that are graduate programs or medical schools is an outstanding accomplishment as these future scientists will most certainly provide key research accomplishments in the years to come.

Reportable Outcomes:

The following publications have our students as authors:

Grisanzio C, Werner L, Takeda D, **Awoyemi BC**, et al. Genetic and functional analyses implicate the NUDT11, HNF1B, and SLC22A3 genes in prostate cancer pathogenesis. Proc Natl Acad Sci U S A. 2012 109: 11252-7.

The other students have reported their findings to the University of Iowa faculty, to the faculty and students at Lincoln University, at national competitions and conferences. The following table demonstrates the placement in the Science Fair Competition held at Lincoln University each fall.

Year	Name of Student	Mentors	Outcome/Prize
2006	Shaan Spence,	Frederick Domann and Michael Hitchle	Hon. Mention: Biol. Science; Cat 1
	Briquel Sherman	Sudhanshu Raikwar	2 nd prize: Biol. Science; Cat 2
2007	Elizabeth Okyne,	Melissa Fath, Andrean L. Simons,	1 st : prize Cancer Research
		Douglas R. Spitz	and Safford's cell biology award
	Seme Diallo,	Sabrina Bonde and Nicholas Zavazava	2 nd prize: Cancer research
	Katrina Probherbs,	David Lubaroff and Susan Lutgendorf	2 nd prize: Cancer research
	Patrick Ndungu,	Dev. Karan and David Lubaroff	1 st prize: Cancer therapy & prevention; Best overall poster
	Caroline O Dias,	Elizabeth Chrischilles, and Gerry Rushton	3 rd prize : Cancer therapy & prevention
2008	Theon Francis	Todd Rouse, ME Nasr and EH Field	2 nd prize: excellent poster: all categories
	Stacy-ann Wright	Susan Wiechert & JR Bickenbach	Hon. Mention:
			all categories
	Michelle Gray	Trent Waugh & David Lubaroff	3 rd prize: great poster:
			all categories
	Julia Greenfield	CJ Weydert, Allison Esser & MD Henry	2 nd prize: excellent poster: all categories
	Bukola Fatunmbi	Jessica Graham, Cait Lemke & Aliasger	Hon. Mention:
	1	Salem	all categories
2009	Stephen C. Sangster	Adam J. Case and Frederick E. Domann	2 nd prize: Cell/Mol/Env Sci.
	Kaylene K. Baugh	Van S. Tompkins and Siegfried Janz	3 rd prize: Cancer Biology
2010	Theresa Akede	Yi Luo	Hon. Mention Cell/Mol/Env Sci.
	Stephanie Rand	Suresh Veeramani and George Weiner	2 nd prize: Cell/Mol/Env Sci.
	Danielle Mcknight	Douglas R. Spitz and Melissa A. Fath	3 rd prize: Cancer Prev & Treatment
	Stephen C. Sangster	Timothy R. Rosean, Van S. Tompkins, and Siegfried Janz	2 nd prize: Cancer Prev & Treatment
	Kaylene Baugh	Erica Flanders, James Brown, Keith Crouch and David Lubaroff	Hon. Mention: Cancer biology: Diagnosis
	Nakita Brown	Chris Kowalski, Douglas R. Spitz and Andrean L. Simons	2 nd prize: Cancer biology: Diagnosis
2011	Jhoneil Cooper	Samuel Lifton and Siegfried Janz	Hon Mention: Cat A
	Nathaniel J. Sangster	Douglas R. Spitz and Frederick Domann	3 rd prize: Cat A
	Patrick Ihejirika	Kyle C. Kloepping, Douglas R. Spitz and Michael K. Schultz	1 st prize: Cat B
	Ayanna Raeburn	Arya Sobhakumari and Andrean L. Simons-Burnett	2 nd prize: Cat C
	Candice Marie Lynch	David Lubaroff, and Aliasger Salem	1 st prize: Cat C Best overall poster
	Darah Doubt- Swinton	David Lubaroff	Hon mention: Cat C
	Jodi-Ann Foster	Suresh Veeramani and George Weine	3 rd . prize: Cat C

Conclusion

This award was highly successful as evidenced by the amount of work accomplished by each student and by their motivation to continue in a science career. The PI applied, and received funding, for a second HBCU training grant that will enable us to accept additional students for the next 3 years, thus increasing the number of African American scientists in the area of prostate cancer.

Appendices: Brochures for 2010, 2011, and 2012





2010 Prostate Cancer Research Summer Training Program

A Collaboration Between the University of Iowa and Lincoln University of Pennsylvania



Students in the 2009 Program

Summary of Program: The partnership of the University of Iowa and Lincoln University is designed to provide an outstanding atmosphere to train undergraduate students from Lincoln in prostate cancer research. We propose to have fourteen mentors available for each of the trainees to choose for their summer research project. The mentors are from seven departments and three colleges at the University of Iowa and the prostate cancer research in their laboratories covers a wide a rea of interest. The proposed mentors have extensive training experience at all levels; under graduate, graduate, medical, and postdoctoral.

In addition to the 14 faculty mentors both the University of Iowa and Lincoln University have designated Faculty Advisors for the students. Dr. Paul Heidger serves as the advisor at the University of Iowa and Dr. Derrick Swinton serves as the advisor at Li ncoln University. Both individual s are available for advice a nd assistance throughout the summer and the regular academic year. The faculty members are listed below as well as a brief description of research in the la boratories of each University of Iowa mentor.

At this point in time the program is 8 weeks long, beginning on Monday, June 7, 2010 and ending on Friday, July 30, 2010.

Faculty Advisor at Lincoln University: Derric k Swinton, PhD.; Associate Professor, Department of Analytical Chemistry (610- 932-8300, ext.3470) http://www.lincoln.edu/chemistry/swinton.html

University of Iowa Faculty and Their Research

Director and Research Mentor: David Lubaroff, PhD; Professor, Department of Urology & Director of the Summer Research Program (319-335-8423) http://www.uihealthcare.com/depts/med/urology/urologymds/lubaroff.html

The work in this laboratory concentrates on the area of tumor i mmunology with an emphasis on immunotherapy. We havve constructed microbial vaccines to be used for the investigation of gene and immunotherapy of prostate cancer. Investigations on the ability of immunized animals to produce immune responses to the transgene product induced by the vaccine are underway. Additionally, we are carrying our "translational" research in the form of clinical trials of our adenovirus vaccine in men with prostate cancer. Important in these trials is the safety of the vaccine and its ability to induce anti-tumor immunity. We have recently completed a Phase I clinical trial of the vaccine that demonstrated its safety. Current plans are underway to initiate a therapeutic Phase II trial. Finally, we have b een collaborating on studies of psychosocial effects on immune status in cancer patients.

Faculty Advisor: Paul Heidger, PhD; Professor, Depat. of Anatomy & Cell Biology (319-335-7722) http://www.anatomy.uiowa.edu/pages/directory/faculty/heidger.html

Dr. Heidger will assist in the recruitm ent and evaluation of summer students and will assist in career planning.

Research Mentors

Jackie R. Bickenbach, Ph.D.; Associate Professor, Department of Anatomy & Cell Biolo gy (319-335-6719)

http://www.anatomy.uiowa.edu/pages/directory/faculty/bickenbach.html

The research in the Bicken bach lab involves the understanding of how aging affects keratinocye stem cells and developing molecular mechanisms to dedifferentiate and trans-differentiate skin keratinocytes into cells that behave like stem cells, and how cell migration of keratin ocytes and cancer cells are affected. Previously, the lab identified a subset of basal skin keratino cytes as stem cells. These cell s had multipotent characteristics in the at they can differentiate into various other types of cells and tissues. Currently, they are looking for specific markers for these stem cells, and trying to determine whether they have a ctivated different si gnaling pathways. Of primary interest is whether these stem cells have potential clinical relevance, especially in age-related diseases, particularly cancers. Dr. Bickenbach's lab h as shown that the age of the keratinocyte stem cell has little effect on its multipotent capabilities, and thus could be used in translational or clinical cell-based therapies. In another project, the lab is using specific transcription factors to de-differentiate skin keratinocytes into cells that behave similar to stem cells. This translational project produces cells that can be tested in models of human disease.

Elizabeth Chrischilles, Ph.D.; Professor, Department of Epidemiology (319-384-5009)

http://www.public-health.uiowa.edu/epi/faculty/elizabeth_chrischilles.html

Dr. Chrischilles directs the Health Effectiveness Research Center (HERCe) (www.publichealth.uiowa.edu/herce/), a collab orative research enterprise between the Department of Epidemiology and the Coll ege of Pharmacy at the University of Iowa. HERCe focuses on understanding the reasons for and consequences of treatment variation in clinical practice. It is a center for research, learning, and education that is comprised of epidemiologists, economists, biostatisticians, clinicians, database specialists, geographers, and graduate students from

colleges and departments across campus. Areas of conceptualization expertise include measurement of preventive care and treatments from retrospective data; methodolo gies for addressing treatment selection bias including instrumental variables and direct statistical and design control for confounding; population-based sampling; analysis of complex sample surveys and I ongitudinal data; geographical analysis of healthcare access; data linkage and application of encryption methodologies to maintain confidentiality; and synth esis of d rug information to evaluate medication safety. Examples of HERCe research include recent publications on breast cancer treatments. complications chemotherapy for lymphoma p atients, invasive treatments for acute myo cardial infarction, and an evaluation of the Iowa Medicai d Pharmaceutical Case Management program.

Frederick Domann, PhD.; Professor, Dept. of Radiation Oncology. (319-335-8018) http://www.uihealthcare.com/depts/med/radiationoncology/frrb/faculty/domann.html

The Domann laboratory is predominantly interested in the regulation of gene expre ssion in cancer that does not in volve classical changes in the DNA sequence, but rather is mediated through so-called "epigenetic" events. These include DNA methylation, histone modifications that affect DNA accessibility, and chromatin conformational changes that rend er genes available or unavailable for efficient transcription. Duri ng a typical summer research experience the und ergraduate student would learn how to dev elop and test a sci entific hypothesis related to a fundamental question in cancer research using state of the art te chniques and approaches. Methods learned would include human cell culture, nucleic acid extraction, conventional PCR, reversetranscriptase-PCR to measure m RNA, real-time quantitative PCR, DNA sequencing, DNA methylation analysis, western blotting, enzyme assays, and molecular cloning. The student would be come proficient at the techniques through daily interactions with laboratory staff. In addition, the student would become familiar with the theory behind each technique and interpretation of their laboratory results through twice weekly meetings with Professor Domann. It is the goal of this research experience to allow the student the opportunity to participate in larger ongoing research projects in the lab in a substantive way so that he or she can contribute to a publication

Elizabeth H. Field, M.D.; Professor, Department of Internal Medicine (319-339-7078) http://www.int-med.uiowa.edu/Divisions/Rheumatology/Directory/ElizabethField.html

CD4+CD25+ T regulato ry (Treg) lymphocytes are both beneficial and deleterious to health, maintaining tolerance to autoantigens and alloanti gens on the one hand while preve nting immunity to tumor or pathogens on the other. Because of their dual effect it is important to define their mechanism of action. The Field laboratory utilizes f our dimensional live cell imaging and fluorescent fusion proteins to probe the functional interactions of CD4+CD25+ regulatory cells. Because CD4+CD25+ Treg cell activity require cell:cell contact, IL-2, and CD25, one project define s the dynamic processe s involved in the paracrine delivery of IL-2 to T reg cells. The I ab generates various IL-2 and CD25 fl uorescent fusion proteins and expresses these in live cells to image IL-2 a nd CD25 intra- and intercellular trafficking profiles and dynamic protein:protein interactions between cells. Other projects characterize the dynamic and functional interactions of CD4+CD25+ T regulatory and conventional CD4+ cells or dendritic cells in a model of skin inflammation and a mouse model of HPV cancer.

Thomas Griffith, PhD; Associate Professor, Department of Urology (319 335 7581) http://www.uihealthcare.com/depts/med/urology/urologymds/griffith.html

The research in the Griffith laboratory studies the therapeutic potential of apoptotic cell death in the treatment of cancer. The tumor necrosis family member TRAIL/Apo-2 ligand is a potent inducer of tumor cell apoptosis, but is non-toxic against normal cell and tissues, su ggesting that TRAIL might be administered as an antitumor therapeutic without the side effects seen with other TNF family members, namely TNF and Fa s ligand, a nd traditional chemotherapeutics. Empl oyment of various g ene delivery systems, such as no n-replicative viral vectors, is making it possible to administer genes directly into tumors sites in situ. Using this technology, a re combinant, replication-deficient adenoviral vector encoding the full -length TRAIL cDNA (Ad-TRAIL) was developed in the laboratory as induce tumor cell death. Current experiments are investigating the ability of Ad-TRAIL to activate systemic antit umor immunity. Additional studies are investigating the role of the innat e immune system, specifically neutrophils, in the antitumor response activated by Mycob acterium bovis BCG intravesical therapy for bladder cancer. Recent studies in our laboratory demonstrated that TRAIL is induced by BCG treatment, and TRAIL levels in the urine correlated with effective therapy. Of th leukocytes present in the u rine, neutrophils expressed high levels of TRAIL. In vitro, human peripheral blood neutrophils contain large intracellular stores of functional TRAIL that is released after stimulation with BCG. Current studies are investigating the contributions of neutrophils in BCG

therapy for blad der cancer, and examining the mechanisms behind BCG-induced expression of TRAIL by neutrophils

Michael Henry, PhD; Associate Professor, Department of Physiology & Biophysics (3 19-335-7886)

http://www.physiology.uiowa.edu/faculty/faculty/henry.htm

Research in the Henry laboratory is geared toward understanding the mole cular and cellular biology underlying the sp read of can cer cells from the prostate to other vital organs such as bone, liver and lung. They have developed animal models of prostate cancer met astasis that employ bioluminescence imaging to visualize metastatic cancer cells in living animals. A summer research project would be to enginee r and characterize a prostate cancer cell line for expression of the firefly luciferase gene so that it might be used in our animal models.

Siegfried Janz, MD; Professor, Department of Pathology (319-384-2869)

http://www.healthcare.uiowa.edu/pathology/site/faculty/janz/janz.html

Siegfried Janz' primary research interest concerns mouse models of hu man B cell a nd plasma cell neoplasms that are in duced by the deregulated expression of the cell ular oncogene MYC (c-myc). His laboratory has recently generated gene-insertion mice that mimic three diff erent states of the human genetic alterations. He is now developing genetic methods for the detection of the homologous Mycactivating translocations in mice. As leader of the Cancer Genetics and Computational Biology Program at the Holden Comprehensive Cancer Center, he is also a ctively engaged in research on human blood cancers.

Yi Luo, MD, PhD; Assistant Professor, Department of Urology (319-335-9835) http://www.uihealthcare.com/depts/med/urology/urolo

gymds/luo.html

A major research project in our la boratory is to develop a novel therapeutic strategy to cope with the limitations of the current modalities for prostate cancer treatment. We will use prostate-specifi c antigen (PSA), a protei n known to be aberran tly expressed in prostate cancer, as a target for immunotherapy of prostate cancer. In fact, PSA has been demonstrated to be a useful immunotherapeutic target in clinical trials as well as in animal models. In addition, PSA has al so been demonstrated to be antigenic and capable of inducing specific immune responses in both humans and mice. However, up to

date. all c urrently available PSA-targeted immunotherapies have only demonstrated limited antitumor effects. To improve this immunotherapeutic approach, we will use bot h bacillus Calmette-Guérin (BCG, a bacterial vaccine strain) and adenovirus (Ad, a replication-defective strain) to deliver PSA for animal immunization. Both BCG and Ad microbes have been demonstrated to be safe and effective for antigen delivery in humans and mice. Since the se two microbes are kn own to be different in thei infectious modes and host anti-infection responses, rationally combined use of B CG and A d recombinants for vacci nation will provide a synergistic/complementary immune induction and thus likely result in enhanced antitumor immunity. Indeed, we have previously observed a robust induction of PSA-specific T cell res ponses by vaccination with combined B CG-PSA (primer vaccine) and Ad-PSA (boos ter vaccine) in mic e. In this study, we will further ev aluate the effects of this vaccination method on preventing or treatin g experimental prostate tumors. The objective of this study is to provide a proof of principle that enhanced antitumor immunity can be achieved by combine d vaccination with BCG and Ad recombinants.

Aliasger K. Salem, PhD; Assistant Professor, Division of Pharmaceutics, College of Pharmacy (319-335-8810)

http://www.pharmacy.uiowa.edu/pharmaceutics/people/Salem.htm

Dr. Salem's research interests are primarily focused on self-assembling systems, the rational design of novel drug and gene delivery systems and on the development of sophi sticated scaffolds for tissu especific regeneration. In tissue engineering, Dr. laboratory applies microfabrication Salem's techniques to novel biomaterials to provide spatial control over tissue for mation and to integrate minimally invasive scaffold delivery strategies. In drug/gene delivery, he is currently exploring the synergistic application of degradable particle technology, CpG oli gonucleotides and heat sh ock proteins for generating sustained immunotherapeutic responses against cancer. Dr. Sale m's laboratory also collaborates with Dr. Lubaroff on the use o microparticles in association with cancer vaccines fro the induction of strong anti-tumor immune responses and tumor destruction.

Michael Schultz, PhD; Faculty Associate, Department of Internal Medicine (319-356-4159) http://www.int-med.uiowa.edu/Divisions/Cardiology/Directory/MichaelSchultz.html

Dr. Schultz's laboratory is interested in exploring cellsurface protein expression (e.g., G-coupled protein receptors) that is amplified in specific cancer cell lines and developing peptide- and RNA-aptamerbased molecular targeting mechanisms for delivering radionuclides specifically to the site of cancerous tissue in the body. Examples of Dr. Schultz's include the development of no vel research radiolabeled peptide-analogs of neuropeptide Y (NPY) that are designed to bind with high affinity to neuropeptide Y subtype 2 re ceptors (Y2). In a second example of Schultz laboratory research, an ribonucleic acid (RNA) compou nd (known as an aptamer) has been synthesized that binds tightly to a cell surface protein receptor (referred to as PSMA) whose expression is amplified on the surface of prostate cancer cells relative to normal cells. Through the development of a novel chelator derivative, Dr. Schultz and colleagues are able to radiola bel the aptamer for imaging by PET. These exciting imagin g agents serve not only as high resolution probes for evaluating the location and extent of disease, but also pave the way for the d evelopment of molecularly-guided therapeutic agents that hold promise in the development of curative approaches to these enigmatic cancers.

Douglas Spitz, PhD; Professor, Department of Radiation Oncology (319-335-8001) http://www.uihealthcare.com/depts/med/radiationonc ology/frrb/faculty/spitz.html

Research in the Spitz lab oratory is concentrated on the role of free radicals and oxidative events in cancers. For example, combinations of inhibitors of glucose metabolism, 2-deoxy-D-glucose (2-DG), and of hydroperoxide detoxification, dehydroisoandrosterone (DHEA) and L-buthionine sulfoximine (BSO), have been shown to be effective in killing hum an tumor cells via oxidative stress. 2-DG has also been shown to increase radiosensitivity in human can cer cells both in vitro and in vivo . These results have led us to test the ability of 20 mM 2-DG + 300 μ M DHEA + 1 mM BS O to induce radiosensitization following exposure to 4 Gy ionizing ra diation. Clonogenic survival was used as the parameter indicative of cytotoxicity. Prostate cancer cells (PC-3) treated with 2-DG or DHEA alone as well as the combinations of 2-DG + DHEA, 2-DG + BSO, DHEA + BSO, or 2-DG + DHEA + BSO all demonstrated so me degree of radiosensitization, and the effect was most pronounced in the group treated with 2-DG + DHEA + BSO. relative to the other combi nations (< 2% survival in the 2-DG + DHEA + BSO group versu s > 5% with oth er agents). In anothe r human prostate cancer cell line, DU145, 2-DG + DHE A + BSO also resulted in substantially enhanced radiosensitization when compared to any of the oth er combinations. These results supp ort the hypothesis that the combining inhibitors of glucose metabolism with inhibitors of hydroperoxide detoxification increases radiation sensitivity in human cancer cells.

George Weiner, MD; Professor, Department of Internal Medicine and Director, Holden Comprehensive Cancer Center (319-353-8620) http://www.int-med.uiowa.edu/Divisions/HemOnc/Directory/George Weiner.html

The laboratory of Dr. G eorge Weiner focuses on exploring methods to enhance the efficacy of monoclonal antibody therapy of cancer. Preclini cal and clinical studies are exploring the relative role of various effector cells in antibody dependent cellular cytotoxicity, how complement impacts on the efficacy of monoclonal antibody therapy and how therapy can be improved. Dr. Wei ner's laboratory is also evaluating the use of other immunotherapy agents such immunostimulatory CpG oligodeoxynucleotides (CpG O DN). He works closely with Dr. Brian Link who le ads the cli nical research aspects of their collaborative r esearch program. Dr. Weiner is the Director of the University of Iowa Holden Comprehensive Cancer Center, and of the Iowa/ Mayo Clinic Speci alized Program of Research Excellence (SPORE) in lymphoma. He is also the principal investigator of additional research grants from the National Cancer Institute and the Leukemia and Lymphoma Society in the field of immunotherapy of cancer.

Nicholas Zavazava, MD, PhD; Professor, Department of Internal Medicine (319-384-6577) http://www.healthcare.uiowa.edu/InternalMedicine/Divisions/Allergy/Directory/NicholasZavazava.html

Research in the Zavazav a laboratory is devoted to the characterization of primate embryonic stem cells. The motivation for this emphasis is that can cer appears to originate from cancer stem cells. These cancer cells have not been well characterized, but appear to share basic characteristics with embryonic stem cells, for example the property of uncontrolled growth. Characterization of these cells will some day lead to better treatment of cancer. Our laboratory is interested in characterizing primate embryonic stem cells and understanding their properties that allow self renewal and immune evasion. Further, the lab is interested in differentiating these cells in vitro into T cells that could be used for the treatment of cancer in the in vivo situation.

Research Facilities - The research laboratories of the faculty mentors at the University of Iowa are located on the west side of Iowa City in the Health Sciences Campus. The facilities include the Medical Laboratories, Bowen Sciences Building, Pharmacy Building, UI General Hospital, Medical Education and Biomedical Research Facility, and the Veterans Affairs Medical Center. Support for the research is provided by a large number of Shared Core Facilities that include the Gene T ransfer Vector Core, DNA Core, Flow Cytometry Core, to name but a few. For that inclu des laboratory research professional, humane veterinary care is provided by the Animal Care Facilities of the Univ ersity of Iowa and the Veterans Affairs Medical Center.

Opportunities for Learning - Students will have a large number of opportunities to learn about research, prostate cancer, and cancer in general. These include meeting with other members of the HBCU SRT and mentors, joint laboratory meeting s with other investigators collaborating with the mentor, journal clubs, and a six-week course designed to educate the students about prostate cancer, its origins, genetics, epidemiology, and treatment.

Living in Iowa City for the Summer

Housing and Meals - All students will be housed in the Mayflower Residence Hall on the Campus of the University of Iowa. It is convenie ntly located on the northern edge of the campus and is served by the free Cambus transportation system. The livin g quarters are also across the Iowa River from the Iowa City Park

Arrival and Welcome – For the 8 week program, students will be expected to arrive on Saturday, June 5, 2010. Flights by most major airlines are available to the Ceda r Rapids Eastern Iowa Airport (CID). These include American, Delta, Northwest, and United Airlines. A welcoming barbecue will be held at City Park on Sunday, June 6 with members of other summer research programs that include the Iowa Biosciences Advantage, and the Student Summer Research Opportunities Program.

Activities In and Around Iowa City - There are a number of activities in the Iowa Ci ty Area that students can find duri ng the sum mer research program. These include, but not li mited to, the following:

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lowa City Jazz Festival – A free, three -day jazz concert featuring lo cal, regional, and national jazz groups during the July 4 th celebration. The festiva I

will be held on the Pentacrest on the campus of the University of Iowa.

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Application to the Prog ram - Application forms, distributed with this brochure, must be completed and returned either to Dr. Swinton at Lincoln University or to Dr. Lubaroff at the University of Iowa. The deadline for submission is Mar ch 5, 201 0. A committee composed of Dr. Swinton, Dr. Luba roff, Dr. Heidger and two a dditional faculty from the University of Iowa will meet and make final decisions. Students will be notified of the decisions no later than March 19, 2010 pending prompt receipt of all applications.

Financial Support - The housing, meal, a nd transportation costs will be paid by the program. In addition, each student will be provided a stipend, the amount of which i s currently being ne gotiated with the University of Iowa and Lincoln University.

For additional information please contact one of the following:

David Lubaroff, PhD, Depart ment of Urology, University of Iowa, 375 Newton Road, 3210 MERF, Iowa City, IA 52242; 319-335-8423; david-lubaroff@uiowa.edu

Paul Heidger, PhD, Department of Anatomy & Cell Biology, University of Iowa, 51 Newton Road, Iowa City, IA 52242; 319-335-7722; paul-heidger@uiowa.edu.

Derrick Swinton, PhD, Department of Analytical Chemistry, Lincoln University, 1570 Baltimore Pike, Lincoln University, PA 19352; 610-932-8300, ext. 3470; dswinton@lincoln.edu

Diane Morman, Program Coordinator, Department of Urology, University of Iowa, 375 Newton Road, 3209 MERF, Iowa City, IA 52242; 319-335-8425; dianemorman@uiowa.edu























2011 Prostate Cancer Research Summer Training Program

A Collaboration Between the University of Iowa and Lincoln University of Pennsylvania



Students in the 2010 Program

Summary of Program: The partnership of the University of Iowa and Lincoln University is designed to provide an outstanding atmosphere to train undergraduate students from Lincoln in prostate cancer research. We propose to have fourteen mentors available for each of the trainees to choose for their summer research project. The mentors are from seven departments and three colleges at the University of Iowa and the prostate cancer research in their laboratories covers a wide a rea of interest. The proposed mentors have extensive training experience at all levels; under graduate, graduate, medical, and postdoctoral.

In addition to the eleven faculty mentors both the University of Iowa and Lincoln University have designated Faculty Advisors for the students. Dr. Paul Heidger serves as the advisor at the University of Iowa and Dr. Derrick Swinton serves as the advisor at Li ncoln University. Both individual s are available for advice a nd assistance throughout the summer and the regular academic year. The faculty members are listed below as well as a brief description of research in the la boratories of each University of Iowa mentor.

At this point in time the program is 8 weeks long, beginning on Monday, June 6, 2011 and ending on Friday, July 29, 2011.

Faculty Advisor at Lincoln University: Derric k Swinton, PhD.; Associate Professor, Department of Analytical Chemistry (610- 932-8300, ext.3470) http://www.lincoln.edu/chemistry/swinton.html

University of Iowa Faculty and Their Research

Director and Research Mentor: David Lubaroff, PhD; Professor, Department of Urology & Director of the Summer Research Program (319-335-8423) http://www.uihealthcare.com/depts/med/urology/urologymds/lubaroff.html

The work in this laboratory concentrates on the area of tumor i mmunology with an emphasis on immunotherapy. We havve constructed microbial vaccines to be used for the investigation of gene and immunotherapy of prostate cancer. Investigations on the ability of immunized animals to produce immune responses to the transgene product induced by the vaccine are underway. Additionally, we are carrying our "translational" research in the form of clinical trials of our adenovirus vaccine in men with prostate cancer. Important in these trials is the safety of the vaccine and its ability to induce anti-tumor immunity. We have recently completed a Phase I clinical trial of the vaccine that demonstrated its safety. We have initiated a therapeutic Phase II trial. Finally, we have been collaborating on studies of psychosocial effects on immune status in cancer patients.

Faculty Advisor: Paul Heidger, PhD; Professor, Depat. of Anatomy & Cell Biology (319-335-7722) http://www.anatomy.uiowa.edu/personnel.shtml?id=heidgerp

Dr. Heidger will assist in the recruitm ent and evaluation of summer students and will assist in career planning.

Research Mentors

Jackie R. Bickenbach, Ph.D.; Professor, Department of Anatomy & Cell Biolo gy (319-335-6719)

http://www.anatomy.uiowa.edu/personnel.shtml?id=bickenbachj

The research in the Bicken bach lab involves the understanding of how aging affects keratinocye stem cells and developing molecular mechanisms to dedifferentiate and trans-differentiate skin keratinocytes into cells that behave like stem cells, and how cell migration of keratin ocytes and cancer cells are affected. Previously, the lab identified a subset of basal skin keratino cytes as stem cells. These cell s had multipotent characteristics in that they can differentiate into various other types of cells and tissues. Currently, they are looking for specific markers for these stem cells, and trying to determine whether they have a ctivated different si gnaling pathways. Of primary interest is whether these stem cells have potential clinical relevance, especially in age-related diseases, particularly cancers. Dr. Bickenbach's lab h as shown that the age of the keratinocyte stem cell has little effect on its multipotent capabilities, and thus could be used in translational or clinical cell-based therapies. In another project, the lab is using specific transcription factors to de-differentiate skin keratinocytes into cells that behave similar to stem cells. This translational project produces cells that can be tested in models of human disease.

Elizabeth Chrischilles, Ph.D.; Professor, Department of Epidemiology (319-384-5009)

http://www.public-health.uiowa.edu/faculty-staff/faculty/directory/faculty-detail.asp?emailAddress=e-chrischilles@uiowa.edu

Dr. Chrischilles directs the Health Effectiveness Research Center (HERCe) (www.publichealth.uiowa.edu/herce/), a collab orative research enterprise between the Department of Epidemiology and the Coll ege of Pharmacy at the University of Iowa. HERCe focuses on understanding the reasons for and consequences of treatment variation in clinical practice. It is a center for research, learning, and education that is comprised of epidemiologists, economists, biostatisticians, clinicians, database

specialists, geographers, and graduate students from colleges and departments across campus. Areas of expertise include conceptualization measurement of preventive care and treatments from retrospective data; methodolo gies for addressing treatment selection bias including instrumental variables and direct statistical and design control for confounding; population-based sampling; analysis of complex sample surveys and I ongitudinal data; geographical analysis of healthcare access; data linkage and application of encryption methodologies to maintain confidentiality; and synth esis of d rug information to evaluate medication safety. Examples of HERCe research include recent publications on breast cancer treatments. complications chemotherapy for lymphoma p atients, invasive treatments for acute myo cardial infarction, and an evaluation of the Iowa Medicai d Pharmaceutical Case Management program.

Frederick Domann, PhD.; Professor, Dept. of Radiation Oncology. (319-335-8018) http://www.uiowa.edu/~frrbp/domann_lab.html

The Domann laboratory is predominantly interested in the regulation of gene expre ssion in cancer that does not in volve classical changes in the DNA sequence, but rather is mediated through so-called "epigenetic" events. These include DNA methylation, histone modifications that affect DNA accessibility, and chromatin conformational changes that rend er genes available or unavailable for efficient transcription. Duri ng a typical summer research experience the und ergraduate student would learn how to dev elop and test a sci entific hypothesis related to a fundamental question in cancer research using state of the art te chniques and approaches. Methods learned would include human cell culture, nucleic acid extraction, conventional PCR, reversetranscriptase-PCR to measure m RNA, real-time quantitative PCR, DNA sequencing, DNA methylation analysis, western blotting, enzyme assays, and molecular cloning. The student would be come proficient at the techniques through daily interactions with laboratory staff. In addition, the student would become familiar with the theory behind each technique and interpretation of their laboratory results through twice weekly meetings with Professor Domann. It is the goal of this research experience to allow the student the opportunity to participate in larger ongoing research projects in the lab in a substantive way so that he or she can contribute to a publication

Michael Henry, PhD; Associate Professor, Department of Physiology & Biophysics (3 19-335-7886)

http://www.physiology.uiowa.edu/henry.shtml?menu=1&tab=facultyTab

Research in the Henry laboratory is geared toward understanding the mole cular and cellular biology underlying the sp read of can cer cells from the prostate to other vital organs such as bone, liver and lung. They have developed animal models of prostate cancer met astasis that employ bioluminescence imaging to visualize metastatic cancer cells in living animals. A summer research project would be to enginee rand characterize a prostate cancer cell line for expression of the firefly luciferase gene so that it might be used in our animal models.

Siegfried Janz, MD; Professor, Department of Pathology (319-384-2869) http://www.healthcare.uiowa.edu/pathology/site/facult

y/janz/janz.html

Siegfried Janz' primary research interest co ncerns mouse models of hu man B cell a nd plasma cell neoplasms that are in duced by the deregulated expression of the cell ular oncogene MYC (c-myc). His laboratory has recently generated gene-insertion mice that mimic three diff erent states of the human genetic alterations. He is now developing genetic methods for the detection of the homologous Mycactivating translocations in mice. As leader of the Cancer Genetics and Computational Biology Program at the Holden Comprehensive Cancer Center, he is also a ctively engaged in research on human blood cancers.

Yi Luo, MD, PhD; Associate Professor, Department of Urology (319-335-9835)

http://www.uihealthcare.com/depts/med/urology/urologymds/luo.html

A major research project in our la boratory is to develop a novel therapeutic strategy to cope with the limitations of the current modalities for prostate cancer treatment. We will use prostate-specifi c antigen (PSA), a protein known to be aberran tly expressed in prostate cancer, as a target for immunotherapy of prostate cancer. In fact, PSA has been demonstrated to be a useful immunotherapeutic target in clinical trials as well as in animal models. In addition, PSA has al so been demonstrated to be antigenic and capable of inducing specific immune responses in both humans and mice. However, up to available PSA-targeted date. all c urrently immunotherapies have only demonstrated limited antitumor effects. To improve this immunotherapeutic approach, we will use bot h bacillus Calmette-Guérin

(BCG, a bacterial vaccine strain) and adenovirus (Ad. a replication-defective strain) to deliver PSA for animal immunization. Both BCG and Ad microbes have been demonstrated to be safe and effective for antigen delivery in humans and mice. Since the se two microbes are kn own to be different in their infectious modes and host anti-infection responses, rationally combined use of B CG and A d recombinants for vacci nation will provide synergistic/complementary immune induction and thus likely result in enhanced antitumor immunity. Indeed, we have previously observed a robust induction of PSA-specific T cell res ponses by vaccination with combined B CG-PSA (primer vaccine) and Ad-PSA (boos ter vaccine) in mic e. In this study, we will further ev aluate the effects of this vaccination method on preventing or treatin g experimental prostate tumors. The objective of this study is to provide a proof of principle that enhanced antitumor immunity can be achieved by combine d vaccination with BCG and Ad recombinants.

Aliasger K. Salem, PhD; Associate Professor, Division of Pharmaceutics, College of Pharmacy (319-335-8810)

http://www.pharmacy.uiowa.edu/pharmaceutics/people/Salem.htm

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Michael Schultz, PhD; Assistant Professor, Department of Radiology (319-356-4159) http://www.medicine.uiowa.edu/Radiology/faculty-staff/faculty/schultz-michael.html

Dr. Schultz's laboratory is interested in exploring cell-surface protein expression (e.g., G-coupled protein receptors) that is amplified in specific cancer cell lines and developing peptide- and RNA-aptamer-based molecular targeting mechanisms for delivering radionuclides specifically to the site of cancerous tissue in the body. Examples of Dr. Schultz's research include the development of no vel

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Douglas Spitz, PhD; Professor, Department of Radiation Oncology (319-335-8001) http://www.uiowa.edu/~frrbp/spitz_lab.html

Research in the Spitz lab oratory is concentrated on the role of free radicals and oxidative events in cancers. For example, combinations of inhibitors of glucose metabolism, 2-deoxy-D-glucose (2-DG), and of hydroperoxide detoxification, dehydroisoandrosterone (DHEA) and L-buthionine sulfoximine (BSO), have been shown to be effective in killing hum an tumor cells via oxidative stress. 2-DG has also been shown to increase radiosensitivity in human can cer cells both in vitro and in vivo . These results have led us to test the ability of 20 mM 2-DG + 300 μ M DHEA + 1 mM BS O to induce radiosensitization following exposure to 4 Gy ionizing ra diation. Clonogenic survival was used as the parameter indicative of cytotoxicity. Prostate cancer cells (PC-3) treated with 2-DG or DHEA alone as well as the combinations of 2-DG + DHEA, 2-DG + BSO, DHEA + BSO, or 2-DG + DHEA + BSO all demonstrated so me degree of radiosensitization, and the effect was m ost pronounced in the group treated with 2-DG + DHEA + BSO. relative to the other combi nations (< 2% survival in the 2-DG + DHEA + BSO group versu s > 5% with oth er agents). In anothe r human prostate cancer cell line, DU145, 2-DG + DHE A + BSO also resulted in substantially enhanced radiosensitization when compared to any of the other combinations. These results supp ort the hypothesis that the combining inhibitors of glucose metabolism with inhibitors of hydroperoxide detoxification increases radiation sensitivity in human cancer cells.

George Weiner, MD; Professor, Department of Internal Medicine and Director, Holden Comprehensive Cancer Center (319-353-8620) http://www.healthcare.uiowa.edu/Labs/Weiner/

The laboratory of Dr. G eorge Weiner focuses on exploring methods to enhance the efficacy of monoclonal antibody therapy of cancer. Preclini cal and clinical studies are exploring the relative role of various effector cells in antibody dependent cellular cytotoxicity, how complement impacts on the efficacy of monoclonal antibody therapy and how therapy can be improved. Dr. Wei ner's laboratory is also evaluating the use of other immunotherapy agents immunostimulatory such as oligodeoxynucleotides (CpG O DN). He works closely with Dr. Brian Link who le ads the cli nical research aspects of their collaborative r esearch program. Dr. Weiner is the Director of the University of Iowa Holden Comprehensive Cancer Center, and of the Iowa/ Mayo Clinic Speci alized Program of Research Excellence (SPORE) in lymphoma. He is also the principal investigator of additional research grants from the National Cancer Institute and the Leukemia and Lymphoma Society in the field of immunotherapy of cancer.

Michael Wright, PhD; Assistant Professor, Department of Mole cular Physiology & Biophysi cs (319-384-1764)

http://www.physiology.uiowa.edu/wright.shtml?menu =1&tab=facultyTab

The Wright Labo ratory is focuse d on defining the composition, activity, and overall cellular function of protein complexes in higher organisms. We utilize quantitative mass spectrometry as a platform to study protein network dynamics in mo del experimental systems. One of the major projects is the mapping of androgen receptor signaling networks in and rogen receptor-related diseases. We are delineatin androgen signaling cascades in hormone-responsive systems with the goal of understanding how aberrant androgen receptor (AR) signaling contributes to the development and p rogression of the AR-relat ed diseases in human prostate cancer. Another project attempts to define molecular biomarkers in androgen receptor-related diseases. This area involves the identification of protein biomarkers in clinical tissue samples of prostate cancer.. We are using both directed and targeted mass spectrometry workflows to identify and quantify tissue biomarkers in radical prostatectomy samples. The goal of this research is to characterize biomarkers to indolent (e.g. organconfined) and lethal (e.g. metastatic) forms of CaP. These studies have the potential to define novel diagnostic, prognostic, and therapeutic biomarkers in the management and treatment of high-risk, organconfined CaP and early-stage, meta static CaP. We are also developing better prote omic workflows to validate tissue bi omarkers in plasma and serum using mass spectrometry-based assays.

Nicholas Zavazava, MD, PhD; Professor, Department of Internal Medicine (319-384-6577) http://www.int-med.uiowa.edu/Divisions/Immunology/Directory/NicholasZavazava.html

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Derrick Swinton, PhD, Department of Analytical Chemistry, Lincoln University, 1570 Baltimore Pike, Lincoln University, PA 19352; 610-932-8300, ext. 3470; dswinton@lincoln.edu

Diane Morman, Program Coordinator, Department of Urology, University of Iowa, 375 Newton Road, 3209 MERF, Iowa City, IA 52242; 319-335-8425; dianemorman@uiowa.edu

















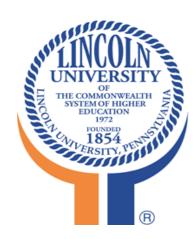












2012 Prostate Cancer Research Summer Training Program

A Collaboration Between the University of Iowa and Lincoln University of Pennsylvania



Students in the 2011 Program

Summary of Program: The partnership of the University of Iowa and Lincoln University is designed to provide an outstanding atmosphere to train undergraduate students from Lincoln in prostate cancer research. We propose to have fifteen mentors available for each of the trainees to choose from for their summer research project. The mentors are from seven departments and three colleges at the University of Iowa and the prostate cancer research in their laboratories covers a wide a rea of interest. The proposed mentors have extensive training experience at all levels; undergraduate, graduate, medical, and postdoctoral.

In addition to the fifteen faculty mento rs both the University of Iowa and Lincoln University have designated Faculty Advisors for the students. Dr. Paul Heidger serves as the advisor at the University of Iowa and Dr. Derrick Swinton serves as the advisor at Li ncoln University. Both individual s are available for advice a nd assistance throughout the summer and the regular academic year. The faculty members are listed below as well as a brief description of research in the la boratories of each University of Iowa mentor.

At this point in time the program is 8 weeks long, beginning on Monday, June 4, 2012 and ending on Friday, July 27, 2012.

Faculty Advisor at Lincoln University: Derric k Swinton, PhD.; Associate Professor, Department of Analytical Chemistry (610- 932-8300, ext.3470) http://www.lincoln.edu/chemistry/swinton.html

University of Iowa Faculty and Their Research

Director and Research Mentor: David Lubaroff, PhD; Professor, Department of Urology & Director of the Summer Research Program (319-335-8423) http://www.uihealthcare.com/depts/med/urology/urologymds/lubaroff.html

The work in this laboratory concentrates on the area of tumor i mmunology with an emphasis on immunotherapy. We havve constructed microbial vaccines to be used for the investigation of gene and immunotherapy of prostate cancer. Investigations on the ability of immunized animals to produce immune responses to the transgene product induced by the vaccine are underway. Additionally, we are carrying our "translational" research in the form of clinical trials of our adenovirus vaccine in men with prostate cancer. Important in these trials is the safety of the vaccine and its ability to induce anti-tumor immunity. We have recently completed a Phase I clinical trial of the vaccine that demonstrated its safety. We have initiated a therapeutic Phase II trial. Finally, we have been collaborating on studies of psychosocial effects on immune status in cancer patients.

Faculty Advisor: Paul Heidger, PhD; Professor, Dept. of Anatomy & Cell Biology (319-335-7722) http://www.anatomy.uiowa.edu/personnel.shtml?id=heidgerp

Dr. Heidger will assist in the recruitm ent and evaluation of summer students and will assist students in career planning.

Research Mentors

Elizabeth Chrischilles, PhD; Professor, Department of Epidemiology (319-384-5009)

http://www.public-health.uiowa.edu/faculty-staff/faculty/directory/faculty-detail.asp?emailAddress=e-chrischilles@uiowa.edu

Dr. Chrischilles directs the Health Effectiveness (HERCe) Research Center (www.publichealth.uiowa.edu/herce/), a collab orative research enterprise between the Department of Epidemiology and the Coll ege of Pharmacy at the University of Iowa. HERCe focuses on understanding the reasons for and consequences of treatment variation in clinical practice. It is a center for research, learning, and education that is comprised of epidemiologists, economists, biostatisticians, clinicians, database specialists, geographers, and graduate students from colleges and departments across campus. Areas of conceptualization expertise include measurement of preventive care and treatments from retrospective data; methodolo gies for addressing treatment selection bias including instrumental variables and direct statistical and design control for confounding; population-based sampling; analysis of complex sample surveys and I ongitudinal data: geographical analysis of healthcare access; data linkage and application of encryption methodologies to maintain confidentiality; and synth esis of d rug information to evaluate medication safety. Examples of HERCe research include recent publications on breast cancer treatments. complications of chemotherapy for lymphoma p atients, invasive treatments for acute myo cardial infarction, and an evaluation of the Iowa Medicai d Pharmaceutical Case Management program.

Frederick Domann, PhD; Professor, Dept. of Radiation Oncology. (319-335-8018) http://www.uiowa.edu/~frrbp/domann_lab.html

The Domann laboratory is predominantly interested in the regulation of gene expre ssion in cancer that does not in volve classical changes in the DNA sequence, but rather is mediated through so-called "epigenetic" events. These include DNA methylation, histone modifications that affect DNA accessibility, and chromatin conformational changes that rend er genes available or unavailable for efficient

transcription. Duri ng a typical summer research experience the und ergraduate student would learn how to dev elop and test a sci entific hypothesis related to a fundamental question in cancer research using state of the art te chniques and approaches. Methods learned would include human cell culture, nucleic acid extraction, conventional PCR, reversetranscriptase-PCR to measure m RNA, real-time quantitative PCR, DNA sequencing, DNA methylation analysis, western blotting, enzyme assays, and molecular cloning. The student would be come proficient at the techniques through daily interactions with laboratory staff. In addition, the student would become familiar with the theory behind each technique and interpretation of their laboratory results through twice weekly meetings with Professor Domann. It is the goal of this research experience to allow the student the opportunity to participate in larger ongoing research projects in the lab in a substantive way so that he or she can contribute to a publication

Michael Henry, PhD; Associate Professor, Department of Physiology & Biophysics (3 19-335-7886)

http://www.physiology.uiowa.edu/henry.shtml?menu=1&tab=facultyTab

Research in the Henry laboratory is geared toward understanding the mole cular and cellular biology underlying the sp read of can cer cells from the prostate to other vital organs such as bone, liver and lung. They have developed animal models of prostate cancer met astasis that employ bioluminescence imaging to visualize metastatic cancer cells in living animals. A summer research project would be to enginee reand characterize a prostate cancer cell line for expression of the firefly luciferase gene so that it might be used in our animal models.

Siegfried Janz, MD; Professor, Department of Pathology (319-384-2869)

http://www.healthcare.uiowa.edu/pathology/site/faculty/janz/janz.html

Siegfried Janz' primary research interest co ncerns mouse models of hu man B cell a nd plasma cell neoplasms that are in duced by the deregulated expression of the cell ular oncogene MYC (c-myc). His laboratory has recently generated gene-insertion mice that mimic three diff erent states of the human genetic alterations. He is now developing genetic methods for the detection of the homologous Mycactivating translocations in mice. As leader of the Cancer Genetics and Computational Biology Program at the Holden Comprehensive Cancer Center, he is also a ctively engaged in research on human blood cancers.

Yi Luo, MD, PhD; Associate Professor, Department of Urology (319-335-9835)

http://www.uihealthcare.com/depts/med/urology/urologymds/luo.html

A major research project in our la boratory is to develop a novel therapeutic strategy to cope with the limitations of the current modalities for prostate cancer treatment. We will use prostate-specifi c antigen (PSA), a protein known to be aberran tly expressed in prostate cancer, as a target for immunotherapy of prostate cancer. In fact, PSA has been demonstrated to be a useful immunotherapeutic target in clinical trials as well as in animal models. In addition, PSA has al so been demonstrated to be antigenic and capable of inducing specific immune responses in both humans and mice. However, up to available PSA-targeted all c urrently immunotherapies have only demonstrated limited antitumor effects. To improve this immunotherapeutic approach, we will use bot h bacillus Calmette-Guérin (BCG, a bacterial vaccine strain) and adenovirus (Ad, a replication-defective strain) to deliver PSA for animal immunization. Both BCG and Ad microbes have been demonstrated to be safe and effective for antigen delivery in humans and mice. Since the se two microbes are kn own to be different in their infectious modes and host anti-infection responses. rationally combined use of B CG and A d recombinants for vacci nation will provide a synergistic/complementary immune induction and thus likely result in enhanced antitumor immunity. Indeed, we have previously observed a robust induction of PSA-specific T cell res ponses by vaccination with combined B CG-PSA (primer vaccine) and Ad-PSA (boos ter vaccine) in mic e. In this study, we will further ev aluate the effects of this vaccination method on preventing or treatin g experimental prostate tumors. The objective of this study is to provide a proof of principle that enhanced antitumor immunity can be achieved by combine d vaccination with BCG and Ad recombinants.

Lyse Norian, PhD; Assistant Professor, Department of Urology (319-335-3013)

Dr. Norian's research is focused on developing novel, hiahlv efficacious combinatorial anti-tumor immunotherapies by utilizing pre-clinical models that accurately reflect the diverse physiologies of can cer patients. To accomplish this, it will be imperative to explore and understand interactions between the immune system and other organ systems, particularly as they relate to anti-tumor immunity. Therefore, her investigations focus not only on the inter-relationships between positive and negative immune cell populations, but also on how the microenvironment of different organ sites impacts anti-tumor immunity. Dr. Norian has developed a robust pre-clinical murine model of metastatic renal cell carcinoma, as well as a

model of metastatic prostate cancer, and now plans to use the se models to i nvestigate mechanisms of site-specific tumor-induced dysfunction in dendriti c cells and T cells. She has previou sly worked with a Lincoln student in collab oration with Drs. Luba roff and Salem during the summer of 2010.

Aliasger K. Salem, PhD; Associate Professor, Division of Pharmaceutics, College of Pharmacy (319-335-8810)

http://www.pharmacy.uiowa.edu/pharmaceutics/people/Salem.htm

Dr. Salem's research interests are primarily focused on self-assembling systems, the rational design of novel drug and gene delivery systems and on the development of sophi sticated scaffolds for tissu especific regeneration. In tissue engineering, Dr. Salem's laboratory applies microfabrication techniques to novel biomaterials to provide spatial control over tissue for mation and to integrate minimally invasive scaffold delivery strategies. In drug/gene delivery, he is currently exploring the synergistic application of degradable particle technology. CpG oli gonucleotides and heat sh ock proteins for generating sustained immunotherapeutic responses against cancer. Dr. Sale m's laboratory also collaborates with Dr. Lubaroff on the use o microparticles in association with cancer vaccines fro the induction of strong anti-tumor immune responses and tumor destruction.

Michael Schultz, PhD; Assistant Professor, Department of Radiology (319-356-4159) http://www.medicine.uiowa.edu/Radiology/faculty-staff/faculty/schultz-michael.html

Dr. Schultz's laboratory is interested in exploring cellsurface protein expression (e.g., G-coupled protein receptors) that is amplified in specific cancer cell lines and developing peptide- and RNA-aptamerbased molecular targeting mechanisms for delivering radionuclides specifically to the site of cancerous tissue in the body. Examples of Dr. Schultz's research include the development of no vel radiolabeled peptide-analogs of neuropeptide Y (NPY) that are desig ned to bind with h igh affinity to neuropeptide Y subtype 2 re ceptors (Y2). In a second example of Schultz laboratory research, an ribonucleic acid (RNA) compou nd (known as an aptamer) has been synthesized that binds tightly to a cell surface protein receptor (referred to as PSMA) whose expression is amplified on the surface of prostate cancer cells relative to normal cells. Through the development of a novel chelator derivative, Dr. Schultz and colleagues are able to radiola bel the aptamer for imaging by PET. These exciting imagin g agents serve not only as high resolution probes for evaluating the location and extent of disease, but also pave the way for the d evelopment of molecularly-guided therapeutic agents that hold promise in the development of curative approaches to these enigmatic cancers.

Andrean Simons-Burnett, PhD; Assistant Professor, Department of Radiation Oncology (319-384-4450)

Dr. Simons-Burnett has been an active partici pant in the summer program, p reviously acting as a "big sister" to students while a member of Dr. Dougla's Spitz's laboratory. Her r esearch interests include metabolic oxidative stress in tumors and the role oxidative stress plays in sign al transduction pathways. Her current interests focus on the EGFR/PI3K/Akt pathway signaling and it s involvement with NADPH oxidase activation, glucose metabolism and autop hagy in cancer. Additionally she is interested in inv estigating novel combined modality therapies that target the E GFR/PI3K/Akt pathway and how one can predict sensitivity to these therapies in cancer disease sites.

Elaine Smith, PhD; Professor, Department of Epidemiology, College of Public Heal th (319-384-5014)

Dr. Smith, a recent addit ion to our mentors, is a Professor of Epidemiology in the Colle ge of Publi c Health. She has a number of research interests that will benefit training of our summer students. These include etiology of on cogenic diseases, focused on molecular epidemiology, HPV effects on the development of genital and other cancers; hormones and risk of HPV detection and replication; HPV and perinatal vertical transmission, head and neck cancers and reproductive diseases: HPV and vestibulitis; prostate cancer risk associated with pesticides and sex steroid hormone alterations.

Douglas Spitz, PhD; Professor, Department of Radiation Oncology (319-335-8001) http://www.uiowa.edu/~frrbp/spitz_lab.html

Research in the Spitz lab oratory is concentrated on the role of free radicals and oxidative events in cancers. For example, combinations of inhibitors of glucose metabolism, 2-deoxy-D-glucose (2-DG), and of hydroperoxide detoxification, dehydroisoandrosterone (DHEA) and L-buthionine sulfoximine (BSO), have been shown to be effective in killing hum an tumor cells via oxidative stress. 2-DG has also been shown to increase radiosensitivity in human can cer cells both in vitro and in vivo . These results have led us to test the ability of 20 mM 2-DG + 300 µM DHEA + 1 mM BS O to induce radiosensitization following exposure to 4 Gy ionizing ra diation. Clonogenic survival was used as the parameter indicative of cytotoxicity. Prostate cancer cells (PC-3) treated with 2-DG or DHEA alone as well as the combinations of

2-DG + DHEA, 2-DG + BSO, DHEA + BSO, or 2-DG + DHEA + BSO all demonstrated so me degree of radiosensitization, and the effect was m ost pronounced in the group treated with 2-DG + DHEA + BSO, relative to the other combi nations (< 2% survival in the 2-DG + DHEA + BSO group versu s > 5% with oth er agents). In anothe r human prostate cancer cell line, DU145, 2-DG + DHE A + BSO also resulted in substantially enhanced radiosensitization when compared to a ny of the oth er combinations. These results supp ort the hypothesis that the combining inhibitors of glucose metabolism with inhibitors of hydroperoxide detoxification increases radiation sensitivity in human cancer cells.

George Weiner, MD; Professor, Department of Internal Medicine and Director, Holden Comprehensive Cancer Center (319-353-8620) http://www.healthcare.uiowa.edu/Labs/Weiner/

The laboratory of Dr. G eorge Weiner focuses on exploring methods to enhance the efficacy of monoclonal antibody therapy of cancer. Preclini cal and clinical studies are exploring the relative role of various effector cells in antibody dependent cellular cytotoxicity, how complement impacts on the efficacy of monoclonal antibody therapy and how therapy can be improved. Dr. Wei ner's laboratory is also evaluating the use of other immunotherapy agents such as immunostimulatory CpG oligodeoxynucleotides (CpG O DN). closely with Dr. Brian Link who le ads the cli nical research aspects of their collaborative r esearch program. Dr. Weiner is the Director of the University of Iowa Holden Comprehensive Cancer Center, and of the lowa/ Mayo Clinic Specialized Program of Research Excellence (SPORE) in lymphoma. He is also the principal investigator of additional research grants from the National Cancer Institute and the Leukemia and Lymphoma Society in the field of immunotherapy of cancer.

Michael Wright, PhD; Assistant Professor, Department of Mole cular Physiology & Biophysi cs (319-384-1764)

http://www.physiology.uiowa.edu/wright.shtml?menu =1&tab=facultyTab

The Wright Labo ratory is focuse d on defining the composition, activity, and overall cellul ar function of protein complexes in higher organisms. We utilize quantitative mass spectrometry as a platform to study protein network dynamics in mo del experimental systems. One of the major projects is the mapping of androgen receptor signaling networks in and rogen receptor-related diseases. We are delineatin g androgen signaling cascades in hormone-responsive systems with the goal of understanding how aberrant androgen receptor (AR) signaling contributes to the

development and p rogression of the AR-relat ed diseases in human prostate cancer. Another project attempts to define molecular biomarkers in androgen receptor-related diseases. This area involves the identification of protein biomarkers in clinical tissue samples of prostate cancer. We are using both directed and targeted mass spectrometry workflows to identify and quantify tissue biomarkers in radical prostatectomy samples. The goal of this research is to characterize biomarkers to indolent (e.g. organconfined) and lethal (e.g. metastatic) forms of CaP. These studies have the potential to define novel diagnostic, prognostic, and therapeutic biomarkers in the management and treatment of high-risk, organconfined CaP and early-stage, meta static CaP. We are also developing better prote omic workflows to validate tissue bi omarkers in plasma and serum using mass spectrometry-based assays.

Nicholas Zavazava, MD, PhD ; Professor, Department of Internal Medicine (319-384-6577) http://www.int-med.uiowa.edu/Divisions/Immunology/Directory/NicholasZavazava.html

Research in the Zavazav a laboratory is devoted to the characterization of primate embryonic stem cells. The motivation for this emphasis is that can cer appears to originate from cancer stem cells. The secancer cells have not been well characterized, but appear to share basic characteristics with embryonic stem cells, for example the property of uncontrolled growth. Characterization of these cells will some day lead to better treatment of cancer. Our laboratory is interested in characterizing primate embryonic stem cells and understanding their properties that allow self renewal and immune evasion. Further, the lab is interested in differentiating these cells in vitro into T cells that could be used for the treatment of cancer in the in vivo situation.

Research Facilities - The research laboratories of the faculty mentors at the University of Iowa are located on the west side of Iowa City on the Health Sciences Campus. The facilities include the Medical Laboratories, Bowen Sciences Building, Pharmacy Building, UI General Hospital, Medical Education and Biomedical Research Facility, and the Veterans Affairs Medical Center. Support for the research is provided by a large number of Shared Core Facilities that include the Gene T ransfer Vector Core, DNA Core, Flow Cytometry Core, to name but a few. For that inclu des laboratory research animals. professional, humane veterinary care is provided by the Animal Care Facilities of the Univ ersity of Iowa and the Veterans Affairs Medical Center.

Opportunities for Learning - Students will have a large number of opportunities to learn ab out research, prostate cancer, and cancer in general. These include meeting with other members of the HBCU SRT and mentors, joint laboratory meeting s with other investigators collaborating with the mentor, journal clubs, and a six-wee k course designed to educate the students about prostate cancer, its origins, genetics, epidemiology, and treatment.

Living in Iowa City for the Summer

Housing and Meals - All students will be housed in the Mayflower Residence Hall on the Campus of the University of Iowa. It is convenie ntly located on the northern edge of the campus and is served by the free Cambus transportation system. The Mayflower has kitchen facilities and double air conditioned rooms. The living quarters are also across the Iowa River from the Iowa City Park

Arrival and Welcome – For the 8 week pro gram, students will be expected to arrive on Sunday, June 3, 2012. Flights by most major airlines are available to the Ceda r Rapids Eastern Iowa Airport (CID). These include American, Delta, and United Airlines. A welcoming barbecue will be held on Sunday, June 3rd with membe rs of other su mmer research programs that inclu de the Iowa Biosciences Advantage, and the Student Sum mer Research Opportunities Program.

Activities In and Around Iowa City - There are a number of activities in the Iowa Ci ty Area that students can find duri ng the sum mer research program. These include, but are not limited to, the following:

Friday and Saturday Night Concert Series – Free musical concerts held e ach Friday and Saturda y night from 6:30 to 9:3 0 pm on the downtown Pedestrian Mall.

lowa City Jazz Festival – A free, three -day jazz concert featuring lo cal, regional, and national jazz

groups during the July 4 th celebration. The festiva I will be held on the Pentacrest on the campus of the University of Iowa.

Thursday Night Concerts in Coralville – These musical concerts, held in Morrison Park in the adjacent town of Coralville, IA, are also free and open to the public.

Saturday Night Free Movies Series – This is the newest addition to lowa City's long tradition of free, outdoor family-friendly entertainment that literall y brings our community together. It is held outdoors on the Pentacrest from June through August.

Other Activities – there are a large number of indoor & outdoor activities that can be accessed through the Cities of lowa City and Coralville and the University of lowa. These include exercise facilities (running, tennis, basketball, volleyball, hand ball/racquetball, weights, biking, and swimming), local beaches, and museums (art, natural h istory, and sports). In addition, there are a l arge number of restaurants ranging from fast food to fine dining.

Application to the Prog ram - Application forms, distributed with this brochure, must be completed and returned either to Dr. Swinton at Lincoln University or to Dr. Lubaroff at the University of Iowa. The deadline for submission is Mar ch 2, 201 2. A committee composed of Dr. Swinton, Dr. Luba roff, Dr. Heidger and two a dditional faculty from the University of Iowa will meet and make final decisions. Students will be notified of the decisions no later than March 16, 2012 pending prompt receipt of all applications.

Financial Support - The housing, meal, a nd transportation costs will be paid by the program. In addition, each student will be provided a stipend, the amount of which i s currently being ne gotiated with the University of Iowa and Lincoln University.

For additional information please contact one of the following:

David Lubaroff, PhD, Depart ment of Urology, University of Iowa, 375 Newton Road, 3210 MERF, Iowa City, IA 52242; 319-335-8423; david-lubaroff@uiowa.edu

Paul Heidger, PhD, Department of Anatomy & Cell Biology, University of Iowa, 51 Newton Road, Iowa City, IA 52242; 319-335-7722; paul-heidger@uiowa.edu.

Derrick Swinton, PhD, Department of Analytical Chemistry, Lincoln University, 1570 Baltimore Pike, Lincoln University, PA 19352; 610-932-8300, ext. 3470; dswinton@lincoln.edu

Diane Morman, Program Coordinator, Department of Urology, University of Iowa, 375 Newton Road, 3209 MERF, Iowa City, IA 52242; 319-335-8425; dianemorman@uiowa.edu

















